54203-H-PCT-US.ST25.txt SEQUENCE LISTING



- Kwong, Peter D. Hendrickson, Wayne A. <110> Sodroski, Joseph G. Wyatt, Richard T.
- <120> CRYSTALLOGRAPHY-RELATED METHOD FOR IDENTIFYING POTENTIAL INHIBITORS OF THE CD4-GP120 INTERACTION
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- <150> PCT/US98/23905
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- <150> us 08/976,741
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Cys Ser Ser Asn Ile Thr Gly Leu Leu Thr Arg Asp Gly Gly Asn 275 280 285

Ser Asn Asn Glu Ser Glu Ile Phe Arg Pro Gly Gly Asp Met Arg 290 295 300

Asp Asn Trp Arg Ser Glu Leu Tyr Lys Tyr Lys Val Val Lys Ile Glu 305 310 315 320

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Asp Met Val Asp Gln Met His Gln Asp Val Ile Ser Leu Trp Asp Gln 20 25 30

Ser Leu Lys Pro Cys Val Lys Leu Thr Pro Leu Cys Val Asn Cys Asn 35 40 45

Thr Ser Ala Ile Thr Gln Ala Cys Pro Lys Val Thr Leu Asp Pro Ile 50 55 60

Pro Ile His Tyr Cys Ala Pro Ala Gly Tyr Ala Ile Leu Lys Cys Asn 65 70 75 80

Asn Lys Thr Phe Asn Gly Thr Gly Pro Cys Asn Asn Val Ser Thr Val 85 90 95

Gln Cys Thr His Gly Ile Lys Pro Val Ile Ser Thr Gln Leu Leu 100 105 110

Asn Gly Ser Ile Ala Glu Glu Glu Ile Ile Ile Arg Ser Glu Asn Leu 115 120 125

Thr Asn Asn Ala Lys Ile Ile Ile Val Gln Leu Asn Lys Ser Val Glu 130 140

Ile Asn Cys Ala Tyr Cys Asn Ile Ser Arg Asn Glu Trp Asn Ile Thr 145 150 155 160

Leu Gln Trp Val Arg Glu Lys Leu Lys Arg His Phe Pro Asn Lys Thr 165 170 175

Ile Asn Phe Thr Gln Pro Ser Gly Gly Asp Leu Glu Ile Thr Thr His $180 \hspace{1cm} 185 \hspace{1cm} 190$

Ser Phe Asn Cys Arg Gly Glu Phe Phe Tyr Cys Asn Thr Ser Ser Leu 195 200 205

Phe Asn Ser Ser Asp Asn Asn Ser Thr Ile Ile Thr Leu Pro Cys 210 220

Arg Ile Lys Gln Ile Ile Asn Met Trp Gln Gly Val Gly Arg Ala Met 225 230 235 240

Tyr Ala Pro Pro Ile Lys Gly Lys Ile Thr Cys Arg Ser Asn Ile Thr 245 250 255

Gly Leu Leu Thr Arg Asp Gly Gly Glu Thr Ser Glu Thr Asn Ser 260 265 270

Thr Glu Thr Phe Arg Pro Gly Gly Gly Asp Met Arg Asp Asn Trp Arg 275 280 285

Ser Glu Leu Tyr Lys Tyr Lys Val Val Glu Val Lys 290 295 300

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Tyr Met Val Glu Gln Met Gln Glu Asp Ile Ile Ser Leu Trp Asp Gln 20 25 30

Ser Leu Lys Pro Cys Val Gln Met Thr Phe Leu Cys Val Asn Cys Asn 35 40 45

Ser Thr Thr Ile Thr Gln Ala Cys Pro Lys Val Ser Phe Glu Pro Ile 50 55 60

Pro Ile His Tyr Cys Ala Pro Ala Gly Tyr Ala Ile Phe Lys Cys Asn 65 70 75 80

Ser Thr Glu Phe Asn Gly Thr Gly Thr Cys Arg Asn Ile Thr Val Val 85 90 95

Thr Cys Thr His Gly Ile Arg Pro Thr Val Ser Thr Gln Leu Ile Leu 100 105 110

Asn Gly Thr Leu Ser Lys Gly Lys Ile Arg Met Met Ala Lys Asp Ile 115 120 125

Leu Glu Gly Gly Lys Asn Ile Ile Val Thr Leu Asn Ser Thr Leu Asn 130 140

Met Thr Cys Glu Tyr Cys Lys Tyr Asn Ala Thr Asp Trp Gly Lys Ile 145 150 155 160

Leu Lys Gln Thr Ala Glu Arg Tyr Leu Glu Leu Val Asn Asn Thr Gly
165 170 175

Ser Ile Asn Met Thr Phe Asn His Ser Ser Gly Gly Asp Leu Glu Val 180 185 190

Thr His Leu His Phe Asn Cys His Gly Glu Phe Phe Tyr Cys Asn Thr 195 200 205

Ala Lys Met Phe Asn Tyr Thr Phe Ser Cys Asn Gly Thr Thr Cys Ser 210 215 220

Val Ser Asn Val Ser Gln Gly Asn Asn Gly Thr Leu Pro Cys Lys Leu 225 230 235 240

Arg Gln Val Val Arg Ser Trp Ile Arg Gly Gln Ser Gly Leu Tyr Ala 245 250 255

Pro Pro Ile Lys Gly Asn Leu Thr Cys Met Ser Asn Ile Thr Gly Met $260 \hspace{1cm} 265 \hspace{1cm} 270 \hspace{1cm}$

Ile Leu Gln Met Asp Asn Thr Trp Asn Ser Ser Asn Asn Asn Val Thr 275 280 285

Phe Arg Pro Ile Gly Gly Asp Met Lys Asp Ile Trp Arg Thr Glu Leu 290 295 300

Phe Asn Tyr Lys Val Val Arg Val Lys 305

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Ser Thr Val Thr Ser Ile Ile Ala Asn Ile Asp Trp Gln Asn Asn Asn 260 265 270

Gln Thr Asn Ile Thr Phe Ser Ala Glu Val Ala Glu Leu Tyr Arg Leu 275 280 285

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PRT

Homo sapiens

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Leu Val Lys Gln Ala Glu Ser Asn Ile His Leu Leu Phe Glu Gln Thr 20 25 30

Met Arg Pro Cys Val Lys Leu Ser Pro Ile Cys Ile His Cys Asn Asp 35 40 45

Ser Val Ile Lys Glu Ala Cys Asp Lys Thr Tyr Trp Asp Thr Leu Arg 50 60

Val Arg Tyr Cys Ala Pro Ala Gly Tyr Ala Leu Leu Lys Cys Asn Asp 65 70 75 80

Lys Asp Tyr Arg Gly Phe Ala Pro Lys Cys Lys Asn Val Ser Val Val 85 90 95

His Cys Thr Arg Leu Ile Asn Thr Thr Ile Thr Thr Gly Ile Gly Leu 100 105 110

Asn Gly Ser Arg Ser Glu Asn Arg Thr Glu Ile Trp Gln Lys Gly Gly 115 125

Asn Asp Asn Asp Thr Val Ile Ile Lys Leu Asn Lys Phe Tyr Asn Leu 130 135 140

Thr Val Arg Cys Arg Trp Cys His Phe Gln Gly Asp Trp Lys Gly Ala 145 150 155 160

Trp Lys Glu Val Arg Glu Glu Val Lys Lys Val Lys Asn Leu Thr Glu 165 170 175 Page 7

Val Ser Ile Glu Asn Ile His Leu Arg Arg Ile Trp Gly Asp Pro Glu 180 185 190

Ser Ala Asn Phe Trp Phe Asn Cys Gln Gly Glu Phe Phe Thr Cys Lys 195 200 205

Met Asp Trp Phe Ile Asn Tyr Leu Asn Asn Arg Thr Glu Asp Ala Glu 210 215 220

Gly Thr Asn Arg Thr Cys Asp Lys Gly Lys Pro Gly Pro Gly Pro Cys 235 240

Val Gln Arg Thr Tyr Val Ala Cys His Ile Arg Gln Val Val Asn Asp 245 250 255

Trp Tyr Thr Val Ser Lys Lys Val Tyr Ala Pro Pro Arg Glu Gly His 260 265 270

Leu Glu Cys Asn Ser Ser Cys Thr Ala Leu Tyr Val Ala Ile Asp Tyr 275 280 285

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Thr Glu Val His Asn Val Trp Ala Thr His Ala Cys Val Pro Thr Asp 35 40 45

Pro Asn Pro Gln Glu Val Val Leu Val Asn Val Thr Glu Asn Phe Asn 50 55 60

Met Trp Lys Asn Asp Met Val Glu Gln Met His Glu Asp Ile Ile Ser 65 70 75 80

Leu Trp Asp Gln Ser Leu Lys Pro Cys Val Lys Leu Tyr Pro Leu Cys 85 90 95 Page 8

Val Ser Leu Lys Cys Thr Asp Leu Lys Asn Asp Thr Asn Thr Asn Ser 100 105 110 Ser Ser Gly Glu Met Met Met Glu Lys Gly Glu Ile Lys Asn Cys Ser 115 120 125Phe Asn Ile Ser Thr Ser Ile Arg Gly Lys Val Gln Lys Glu Tyr Ala 130 135 140 Phe Phe Tyr Lys Leu Asp Ile Ile Pro Ile Asp Asn Asp Thr Thr Ser 155 160 Tyr Thr Leu Thr Ser Cys Asn Thr Ser Val Ile Thr Gln Ala Cys Pro 165 170 175 Lys Val Ser Phe Glu Pro Ile Pro Ile His Tyr Cys Ala Pro Ala Gly 180 185 190 Phe Ala Ile Leu Lys Cys Asn Asn Lys Thr Phe Asn Gly Thr Gly Pro 195 200 205 Cys Thr Asn Val Ser Thr Val Gln Cys Thr His Gly Ile His Pro Val 210 215 220 Val Ser Thr Gln Leu Leu Leu Asn Gly Ser Leu Ala Glu Glu Val 225 230 235 240 Val Ile Arg Ser Ala Asn Phe Thr Asp Asn Ala Lys Lys Ile Ile Val 245 250 255 Gln Leu Asn Gln Ser Val Glu Ile Asn Cys Thr Arg Pro Asn Asn Asn 260 265 270 Thr Arg Lys Ser Ile Arg Ile Gln Arg Gly Pro Gly Arg Ala Phe Val 275 280 285 Thr Ile Gly Lys Ile Gly Asn Met Arg Gly Ala His Cys Ile Asn Ser 290 295 300 Arg Ala Lys Trp Asn Asn Thr Leu Lys Gln Ile Ala Ser Lys Leu Arg 305 310 315 320 Glu Gln Phe Gly Asn Asn Lys Thr Ile Ile Phe Lys Gln Ser Ser Gly 325 330 335 Gly Asp Pro Glu Ile Val Thr His Ser Phe Asn Cys Gly Gly Glu Phe Page 9

Phe Tyr Cys Asn Ser Thr Gln Leu Phe Asn Ser Thr Trp Phe Asn Ser 355 360 365

Thr Trp Ser Thr Glu Gly Ser Asn Asn Thr Glu Gly Ser Asp Thr Ile 370 375 380

Thr Leu Pro Cys Arg Ile Lys Gln Phe Ile Asn Met Trp Gln Glu Val 385 390 395 400

Gly Lys Ala Met Tyr Ala Pro Pro Ile Ser Gly Gln Ile Arg Cys Ser 405 410 415

Ser Asn Ile Thr Gly Leu Leu Leu Thr Arg Asp Gly Gly Asn Asn Asn 420 425 430

Asn Glu Ser Glu Ile Phe Arg Pro Gly Gly Gly Asp Met Arg Asp Asn 435 440 445

Trp Arg Ser Glu Leu Tyr Lys Tyr Lys Val Val Lys Ile Glu Pro Leu 450 460

Gly Val Ala Pro Thr Lys Ala Lys Arg Arg Val Val Gln Arg Glu 465 470 475

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<213> Homo sapiens

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Glu Asp Ile Ile Ser Leu Trp Asp Gln Ser Leu Lys Pro Cys Val Lys 20 25 30

Leu Thr Pro Leu Cys Val Gly Ala 35 40

<210> 8

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<212> PRT

<213> Homo sapiens

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Gly Ser Cys Asn Thr Ser Val Ile Thr Gln Ala Cys Pro Lys Val Ser 1 5 10 15 Page 10

Phe Glu Pro Ile His Tyr Cys Ala Pro Ala Gly Phe Ala Ile 20 25 30

Leu Lys Cys Asn Asn Lys Thr Phe Asn Gly Thr Gly Pro Cys Thr Asn 35. 40

Val Ser Thr Val Gln Cys Thr His Gly Ile Arg Pro Val Val Ser Thr 50 60

Gln Leu Leu Leu Asn Gly Ser Leu Ala Glu Glu Glu Val Val Ile Arg 70 75 80

Ser Val Asn Phe Thr Asp Asn Ala Lys Thr Ile Ile Val Gln Leu Asn 85 90 95

Thr Ser Val Glu Ile Asn Cys Thr Gly Ala 100 105

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Gly His Cys Asn Ile Ser Arg Ala Lys Trp Asn Asn Thr Leu Lys Gln 10 15

Ile Ala Ser Lys Leu Arg Glu Gln Phe Gly Asn Asn Lys Thr Ile Ile $20 \hspace{1cm} 25 \hspace{1cm} 30$

Phe Lys Gln Ser Ser Gly Gly Asp Pro Glu Ile Val Thr His Ser Phe 35 40 45

Asn Cys Gly Glu Phe Phe Tyr Cys Asn Ser Thr Gln Leu Phe Asn 50 55 60

Ser Thr Trp Phe

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Met Trp Gln Lys Val Gly Lys Ala Met Tyr Ala Pro Pro Ile Ser Gly 20 25 30

Gln Ile Arg Cys Ser Ser Asn Ile Thr Gly Leu Leu Leu Thr Arg Asp 35 40 45

Gly Gly Asn Ser Asn Asn Glu Ser Glu Ile Phe Arg Pro Gly Gly Gly 50 60

Asp Met Arg Asp Asn Trp Arg Ser Glu Leu Tyr Lys Tyr Lys Val Val 65 70 75 80

Lys Ile Glu

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Thr Ala Ser Gln Lys Lys Ser Ile Gln Phe His Trp Lys Asn Ser Asn 20 25 30

Gln Ile Lys Ile Leu Gly Asn Gln Gly Ser Phe Leu Thr Lys Gly Pro 35 40 45

Ser Lys Leu Asn Asp Arg Ala Asp Ser Arg Arg Ser Leu Trp Asp Gln 50 55 60

Gly Asn Phe Pro Leu Ile Ile Lys Asn Leu Lys Ile Glu Asp Ser Asp 65 70 75 80

Thr Tyr Ile Cys Glu Val Glu Asp Gln Lys Glu Glu Val Gln Leu Leu 85 90 95

Val Phe Gly Leu Thr Ala Asn Ser Asp Thr His Leu Leu Gln Gly Gln
100 105 110

Ser Leu Thr Leu Glu Ser Pro Pro Gly Ser Ser Pro Ser Val 115 120 125

Gln Cys Arg Ser Pro Arg Gly Lys Asn Ile Gln Gly Gly Lys Thr Leu 130 140

Ser Val Ser Gln Leu Glu Leu Gln Asp Ser Gly Thr Trp Thr Cys Thr 145 150 155 160

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Leu Ala Trp Tyr Gln Gln Lys Pro Gly Gln Ala Pro Arg Leu Leu Ile 35 40 45

Tyr Gly Ala Ser Thr Arg Ala Thr Gly Val Pro Ala Arg Phe Ser Gly 50 60

Ser Gly Ser Gly Ala Glu Phe Thr Leu Thr Ile Ser Ser Leu Gln Ser 65 70 75 80

Glu Asp Phe Ala Val Tyr Tyr Cys Gln Gln Tyr Asn Asn Trp Pro Pro 85 90 95

Arg Tyr Thr Phe Gly Gln Gly Thr Arg Leu Glu Ile Lys Arg Thr Val 100 105 110

Ala Ala Pro Ser Val Phe Ile Phe Pro Pro Ser Asp Glu Gln Leu Lys 115 120 125

Ser Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe Tyr Pro Arg 130 135 140

Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln Ser Gly Asn 145 150 155 160

Ser Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser Thr Tyr Ser 165 170 175

Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu Lys His Lys 180 185 190

Val Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser Ser Pro Val Thr 195 200 205

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Ser Val Lys Val Ser Cys Lys Ala Ser Gly Asp Thr Phe Ile Arg Tyr 20 25 30

Ser Phe Thr Trp Val Arg Gln Ala Pro Gly Gln Gly Leu Glu Trp Met 35 40 45

Gly Arg Ile Ile Thr Ile Leu Asp Val Ala His Tyr Ala Pro His Leu 50 55 60

Gln Gly Arg Val Thr Ile Thr Ala Asp Lys Ser Thr Ser Thr Val Tyr 65 70 75 80

Leu Glu Leu Arg Asn Leu Arg Ser Asp Asp Thr Ala Val Tyr Phe Cys 85 90 95

Ala Gly Val Tyr Glu Gly Glu Ala Asp Glu Gly Glu Tyr Asp Asn Asn 100 105 110

Gly Phe Leu Lys His Trp Gly Gln Gly Thr Leu Val Thr Val Thr Ser 115 120 125

Ala Ser Thr Lys Gly Pro Ser Val Phe Pro Leu Ala Pro Ser Ser Lys 130 135 140

Ser Thr Ser Gly Gly Thr Ala Ala Leu Gly Cys Leu Val Lys Asp Tyr 145 150 155 160

Phe Pro Gln Pro Val Thr Val Ser Trp Asn Ser Gly Ala Leu Thr Ser 165 170 175

Gly Val His Thr Phe Pro Ala Val Leu Gln Ser Ser Gly Leu Tyr Ser 180 185 190

Leu Ser Ser Val Val Thr Val Pro Ser Ser Ser Leu Gly Thr Gln Thr 195 200 205

Tyr Ile Cys Asn Val Asn His Lys Pro Ser Asn Thr Lys Val Asp Lys 210 220

Lys Val Glu Pro Lys 225